APS/Blooks 3/36/92

d his

```
161 S PDGF OR PLATELET DERIVED GROWTH FACTOR
L1
L2
           2545 S ENDOTHEL? OR FIBROBLAST
L3
            106 S L1 AND L2
L4
              1 S L3 AND 349
L5
              2 S (36 OR 37 OR 38)(2W)(KD OR KILODALTON?) AND L3
L6
             57 S (36 OR 37 OR 38)(2W)(KD OR KILODALTON?)
L7
             67 S (36,000 OR 37,000 OR 38,000)(W)(MR OR MOLECULAR(W)WEIGHT
OR
             60 S L7 NOT L6
L8
L9
            117 S L6 OR L8
L10
              3 S L9 AND L1
```

=> d ti ab in pddfd ccls 1-3

US PAT NO:

5,051,364 [IMAGE AVAILABLE]

(FILE 'USPAT' ENTERED AT 14:52:45 ON 30 MAR 92)

L10: 1 of 3

TITLE:

Anti-lipocortin-I and anti-lipocortin-II monoclonal

antibodies

ABSTRACT:

The present invention provides monoclonal antibodies which are specific for one but not both of human lipocortin-I and human lipocortin-II, as well as cultures of hybridomas and other types of cells producing such antibodies.

INVENTOR:

Clare M. Isacke, La Jolla, CA Ian S. Trowbridge, San Diego, CA

Tony Hunter, San Diego, CA

DATE ISSUED:

Sep. 24, 1991

DATE FILED:

Dec. 20, 1989

US-CL-CURRENT: 435/240.27; 424/85.8, 85.91; 435/70.21; 530/387, 389

US PAT NO:

4,935,233 [IMAGE AVAILABLE]

L10: 2 of 3

TITLE: Covalently linked polypeptide cell modulators

ABSTRACT:

Described is a new class of polypeptide cell modulators characterized by being composed of two covalently linked cell modulators in a linear polypeptide sequence. Such dual function polypeptides have new and particularly useful activities when the component polypeptide cell modulators are interferons, lymphokines or cytotoxins which act through different and specific cell receptors to initiate complementary biological activities.

INVENTOR:

Leslie D. Rell, Thame, United Kingdom

Keith G. McCullagh, Princes Risborough, United Kingdom

Alan G. Porter, High Wycombe, United Kingdom

DATE ISSUED:

Jun. 19, 1990

DATE FILED: Dec. 2, 1985

US-CL-CURRENT: 424/85.5, 85.6, 85.7; 435/69.51; 530/351

US PAT NO:

4,543,439

L10: 3 of 3

TITLE

Production and use of monoclonal antibodies to

phosphotyrosine-containing proteins

ABSTRACT:

A hybridoma cell line is disclosed that secretes monoclonal antibodies

which serve as a high titer, reproductble, biological reagent useful in biological/medical research for isolating and identifying phosphotyrosine-containing proteins. In addition, the antibodies have potential uses in diagnosis of a variety of diseases, including certain cancers. The antibodies, which have demonstrated affinity for a variety of nolecules containing o-phosphotyrosine residues, were prepared using a synthetic analog, p-azobenzyl phosphonate (ABP) covalently linked to a carrier protein, as the artigen.

A. Raymond Frackelton, Jr., East Providence, RI

lerman M. Elmen, Wahen, Ma Alonzo H. Ross, Bensalen, PA

DATE ISSUED: Nepa May 1909 1975 FILED: 100. 13, 1962

US-CL-CURRENT: 435/70.21, 240.27, 946; 935/92

warn be

DIOSIS Numbers 93013430

DIFFERENTIAL BINDING BIOLOGICAL AND BICCHEMICAL ACTIONS OF RECOMBINANT PDOF AA AN AND BE MOLECULES ON CONNECTIVE TISSUE CELLS

GROTERDORST O R: IGARASHI A: LARSON R: SOMA Y: CHARETTE M

DEF. CELL BIOLOGY AMATOMY, UNIV. MIAMI SCH. MED., P.C. BOX GICSON, MIAMI, FLORIDA BOLDS.

The second secon Full Journal Titles Journal of Cellular Physiclesy · Languages ENGLISH

We have compared the biological and biochemical properties of recombinant PDOF AN, AM, and DB using three types of fibroblastic cells: MIH/3T3, human skin fibroblast, and fetal bovine acrtic smooth muscle. FDGF binding, receptor autophosphorylation, phosphatidyl incuitel hydrolysis, as well as chemotactic and mitogenic responses of the cells were analyzed. Phor-AD and papers showed minilar receptor binding, receptor autophosphorylation, and potent biclogical activity for all three of the cell types tested. In contrast, PDOF-AA was biologically active only for the MIH/3T3 cells in which binding gites for FDOF-AA were abundant, but was inactive for bovine acreic smooth nuscle calls and human skin fibroblasts in which binding sites for PDGF-AA were absent. PDGF-AA could not induce any biochemical changes in the human skin fibroblasts or smooth nuscle cells. Western blot studies with anti-Type .alpha. and .beta. PDOF receptor antibodies indicate that the MINATE collected both by the alpha and Lesta. Templors, whereas the human skin fibroblasts and bovine smooth muscle cells contained levels of .beta. receptors. These results indicate that cells possessing high levels of PDGF .bets. receptors only are capable of responding equally well to either PDGF AS or ES.

The second second

RICEIS Munhers 67012267

restable to the contract of th PROTEIN

KARTHA S; ERADHAM D M; GROTENDORST G R; TOBACK F G DEP. MED., BOX 453, UNIV. CHICAGO, 5841 SOUTH MARYLAND AVE., CHICAGO, and the second of the second o

AN J PHYSICL 255 (4 PART 2), 1900. FOCO-FOCO. CODEN: AJPHA Full Journal Titles American Journal of Physiclosy

Nontransformed monkey kidney cells (BBC-1 line), used as a model for renal epithelium, were assayed for release of platelet-derived growth factor (PDOF)-like proteins. DSC-1 calls continuously released a mitogeric activity for fibroblasts and a chamcattractant activity for smooth muscle cells, 'each of which was inhibited AB-90% by an artihody to human PDCF. A ally probe for the Michain gene (c-sis), but not for the A-chain gene, hybridized to mRNA obtained from growing and quiescent walls, cosis gene axpressing and property production was standied ... the presence of known growth-regulatory nolecules. A secreted BSC-1 cell protein identical

vo transforming growth fact<u>or</u> "beta.2 inhibited DNA synthesis in growing cultures induced mark accumulation of c-sis_mRNA without and corresponding increase in the release of FDGF-like a livity. Adenosine synthesis in quiescent cultures and enhanced diphosphate stimulated DNA both c-cis expression and release of PDGF-like activity. However, growing the quiescent cells did not express the PDGF receptor gene or exhibit a mitogenic response to authentic FDGF. Thus the FDGF-like protein released by these kidney epithelial cells could contribute to growth control by a paracrine mechanism. ?s pdgf(w)like not s2 1829 PDGF

134382 LIKE PDGF(W)LIKE 131 52 PDGF(W)LIKE NOT S2 130 ?s s3 and (endothel? or fibooblast?)

> 130 53 43997 ENDOTHEL? 65910 FIBROBLAST?

61 S3 AND (EMDOTHEL? OR FIBROBLAST?) S4

?s s4 and (3899WWX&aaooramobob) or 386(2w)(kd or kilodalton?) or (36000(2n)molec qular) or ((36000(2n)(molecular or relative or dalton?))

>>>Command is too complex--please simplify ?s s4 and (349 or 36 or 36000)

> 61 54 853 349 66080 36 36000 78

S5 S4 AND (349 OR 36 OR 36000)

?t s5/6/all

53

5/6/1

BIOSIS Number: 93085187 9100187

EFFECTS OF T CELLS ON PLATELET-DERIVED GROWTH FACTOR-LIKE PROTEIN SECRETION FROM ENDOTHELIAL CELLS

5/6/2

BIOSIS Number: 88102657 7179912

RETINAL PIGMENT EPITHELIAL CELLS PRODUCE PDGF-LIKE PROTEINS AND SECRETE THEM INTO THEIR MEDIA ?t s5/7/all

5/7/1

9100187 BIOSIS Number: 93085187

EFFECTS OF T CELLS ON PLATELET-DERIVED GROWTH FACTOR-LIKE PROTEIN SECRETION FROM ENDOTHELIAL CELLS

SHADDY R E HANSEN J C COWLEY C G

100 N. MEDICAL DRIVE, SALT LAKE CITY, UTAH 84113.

J HEART LUNG TRANSPLANT 11 (1 PART 1). 1992. 48-56. CODEN: JHLTE

Language: ENGLISH

of accelerated graft arteriosclerosis cause after transplantation is unknown. To examine whether the interactions of T cells and endothelial cells (ECs) could contribute to the cause of this phenomenon, T cells were co-incubated with human umbilical artery endothelial cells (HUAEC) and human vein endothelial cells (HUVEC) and the resulting conditioned medium supernantant was assayed for the presence of platelet-derived growth factor (FDGF)-like protein. FDGF-like secretion was significantly greater from HUAECs co-incubated with T cells ar T-cell/HUAEC ratios of 30:1 (6.9 .+-. 1.1 fmol/106 ECs) and 10:1 (6.0 "+-" 1.1 fmol/106 ECs) than the combined background secretion from HUAECs

Profilike protein secretion above background levels from HUARCS was significantly greater, however, than from HUVECs cominculated with T cells at TrosliveC vatios of IOni (1.0 .+-. 0.4 fmol/106 ECs) and 10:1 (0.75 .4-. 0.36 fmol/106 ECs) (p (0.05). In four experiments, preincubation of HIMECE with aganna, thierferon induced Hampe antigen expression but actually caused a decrease in T-cell-induced PDGRF-like protein secretion above background levels (3.8 .+-. 0.6 fmol/106 ECs) when compared to nonetimulated MUNECs (4.8 .+-. 8.4 fmol/186 ECs; p (0.85). PODE-like protein secretion was minimal at I hour and increased over time to a. maximum at 24 hours. The conclusion is that T cells are capable of inducing secretion of a very potent mitogen, pdgf-like protein, from endothelial cells. This cell-inducing production of PDGF-like protein appears to be primarily or exclusively a property of arterial endothelial cells.

7179912 EIGETS Number: 66162627

RETISOL PIGNEST RETURNICE CELLS PRODUCE DOGMINE PROTEINS AND SECRETE THEN INTO THEIR MEDIA

CAMPOCHIARO P A; SUGG R; GROTEHDORST G; HJELMELAND L M DEP. OPHTHOLMOL., UNIV. VQ. SCH. MED., CHARLOTTESVILLE, VA. 22900. EXP EYE RES 49 (2). (1989) 217-228. CODEN: EXERN Full Journal Title: Experimental Eye Research

Human retinal pigment epithelial cells at confluence was used condition serum-free Dulbecco's modified Eagle's medium. Conditioned media were exhaustively dialyzed against 0.5 H acetic acid, lyophilized, and aubjected to Western blot analysis, using as primary antibody an IgO fraction prepared from noat anticerum directed against human platelet-derived growth factor. Native platelet-derived growth factor was resolved as a band with Mr of 30 kDa under non-reducing conditions, while bands with Mr of 36-38 kDa and 10.5 kDa were resolved from retinal pigment epithelial cell-conditioned media. Acid extracts of retinal pigment epithelial cells also contained bands at 36-39 kDa and media conditioned for 48 hr exhibited much denser bands than media conditioned for 24 hr. No bands were detected when non-immune goat IgO fractions were substituted for prinary antibody and when conditioned media were prepared from several Numer fibroblast lines in the same manner as those prepared from retinal pinhent epithelial cells, no detectable bands or only a faint shadow at 36 in the presence of ISSSInethionine were loaded on an anti-platelet-derived frowth factor IgG affinity column, sluted, and subjected to Mo-polyacrylanide gel electrophoresis. Hands with Mr slightly less than 36 Da and 18 kDa were visualized by autoradiography, demonstrating that the platelet-dérived growth factor-like proteins in retinal pigment épithelial Fill-conditioned media are newly synthesized. Two fractions eluted from the films also\markedly stimulated fibroblast chemotaxis and incorporation of idelithymidine, both of which were newtralized Afti-platelet-derived growth factor IgO. These data suggest that retinal Agnest epithelial cells in culture produce platelet-derived growth 'Actor-like proteins and secrete them into their media where they are Sapable of stimulating fibroblast chemotaxis and proliferation.

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LOCUS
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           Molecular cloning of a three-immunoglobulinlike-domain form of a
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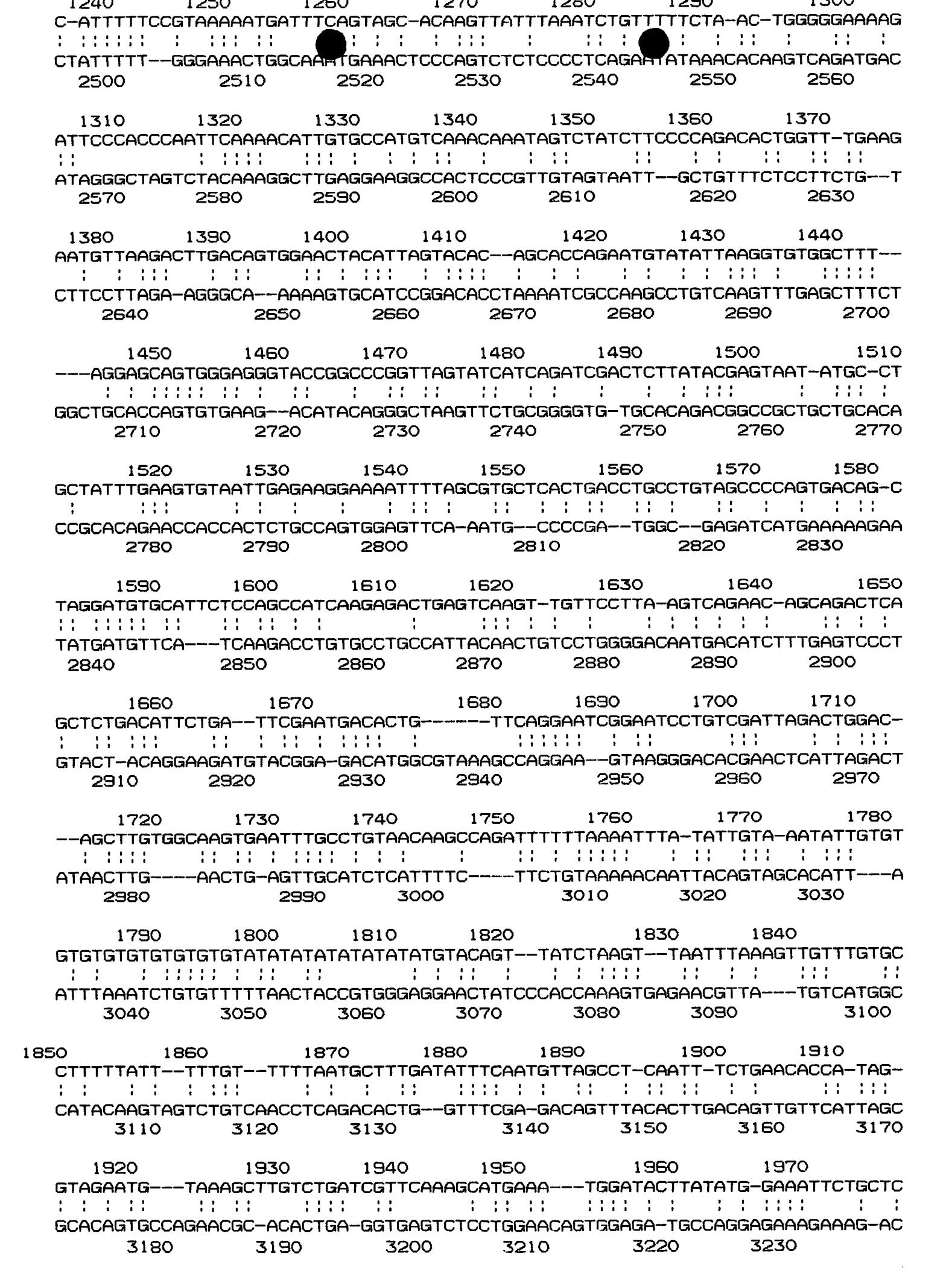
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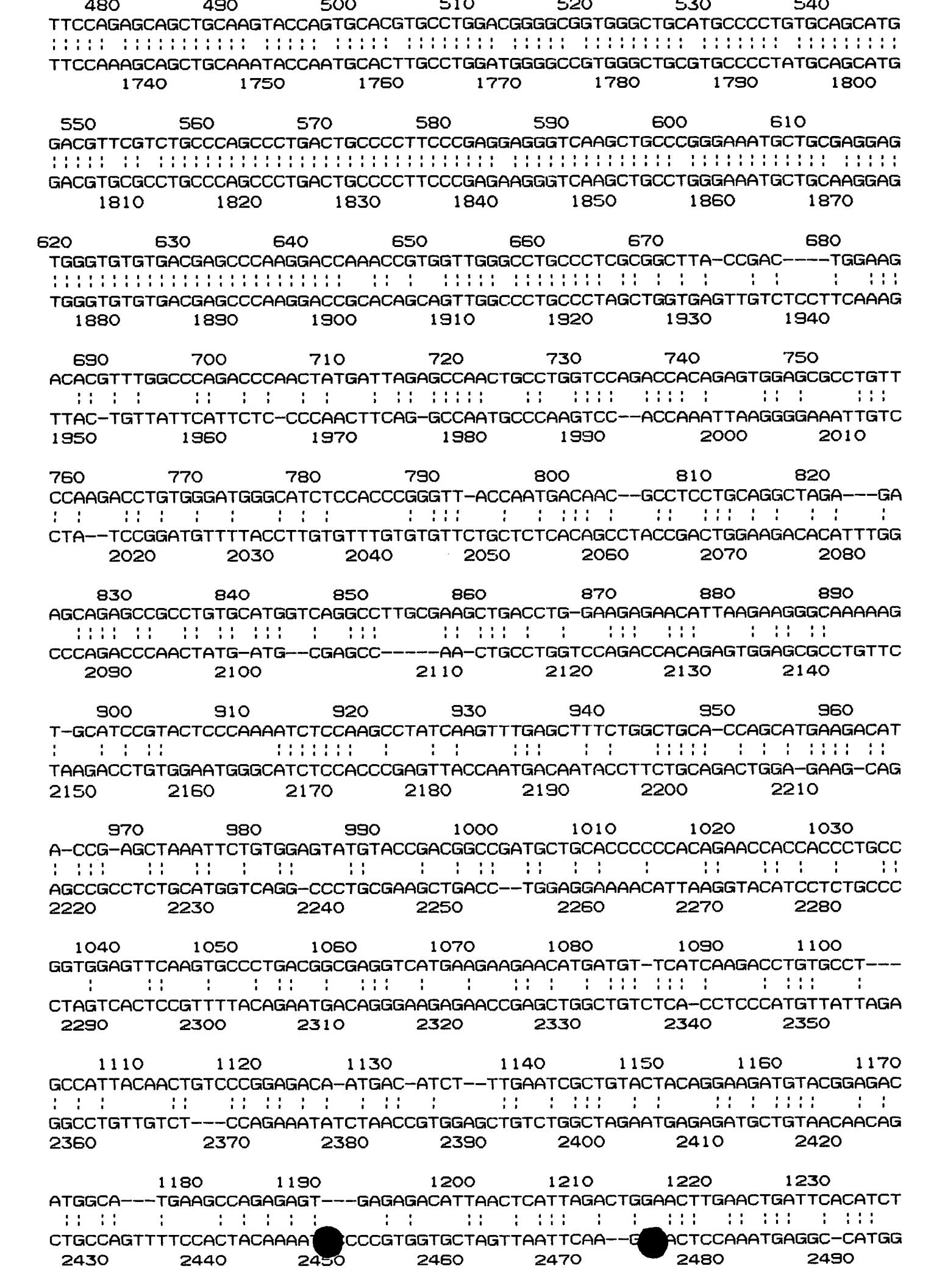
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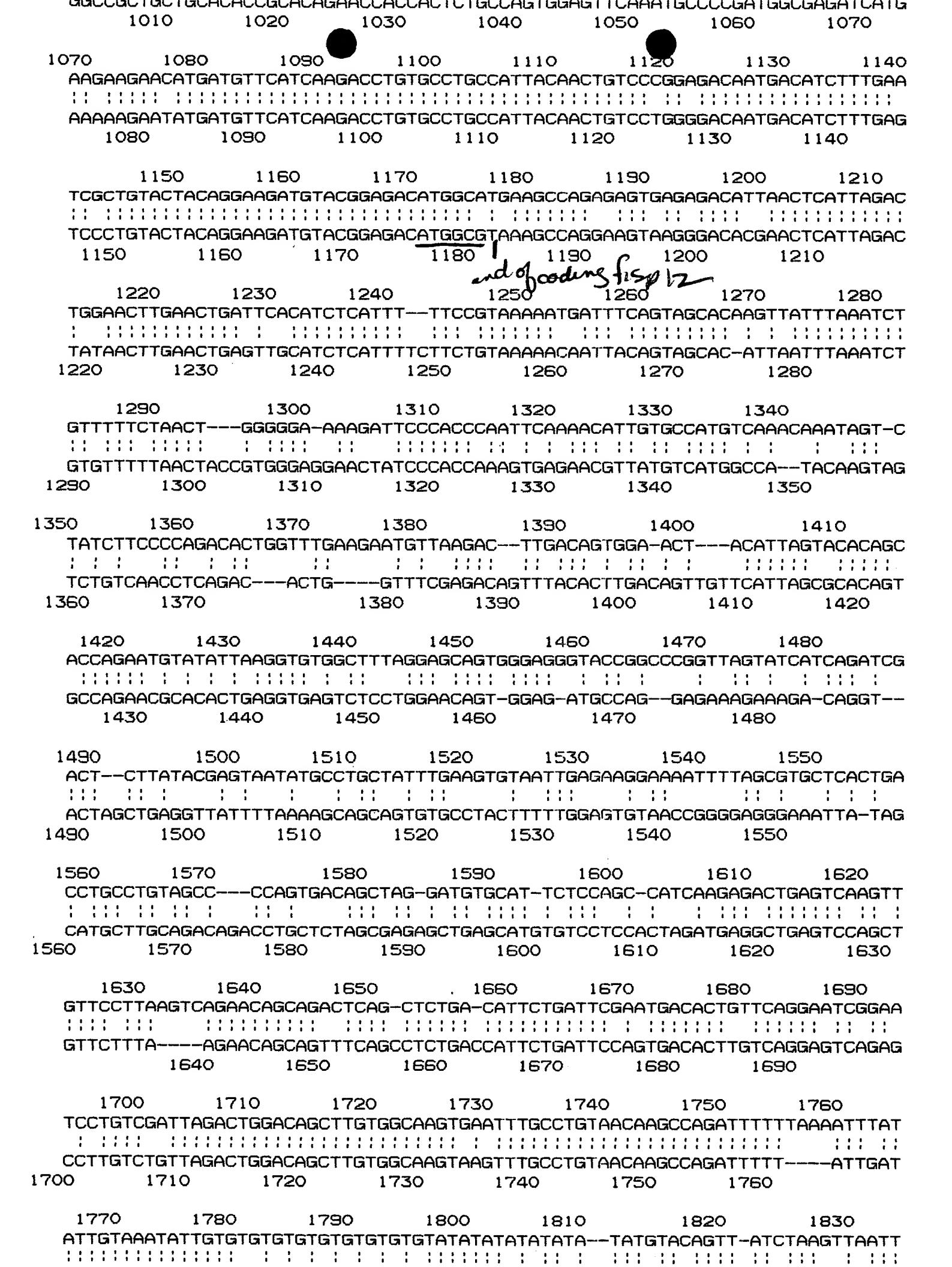
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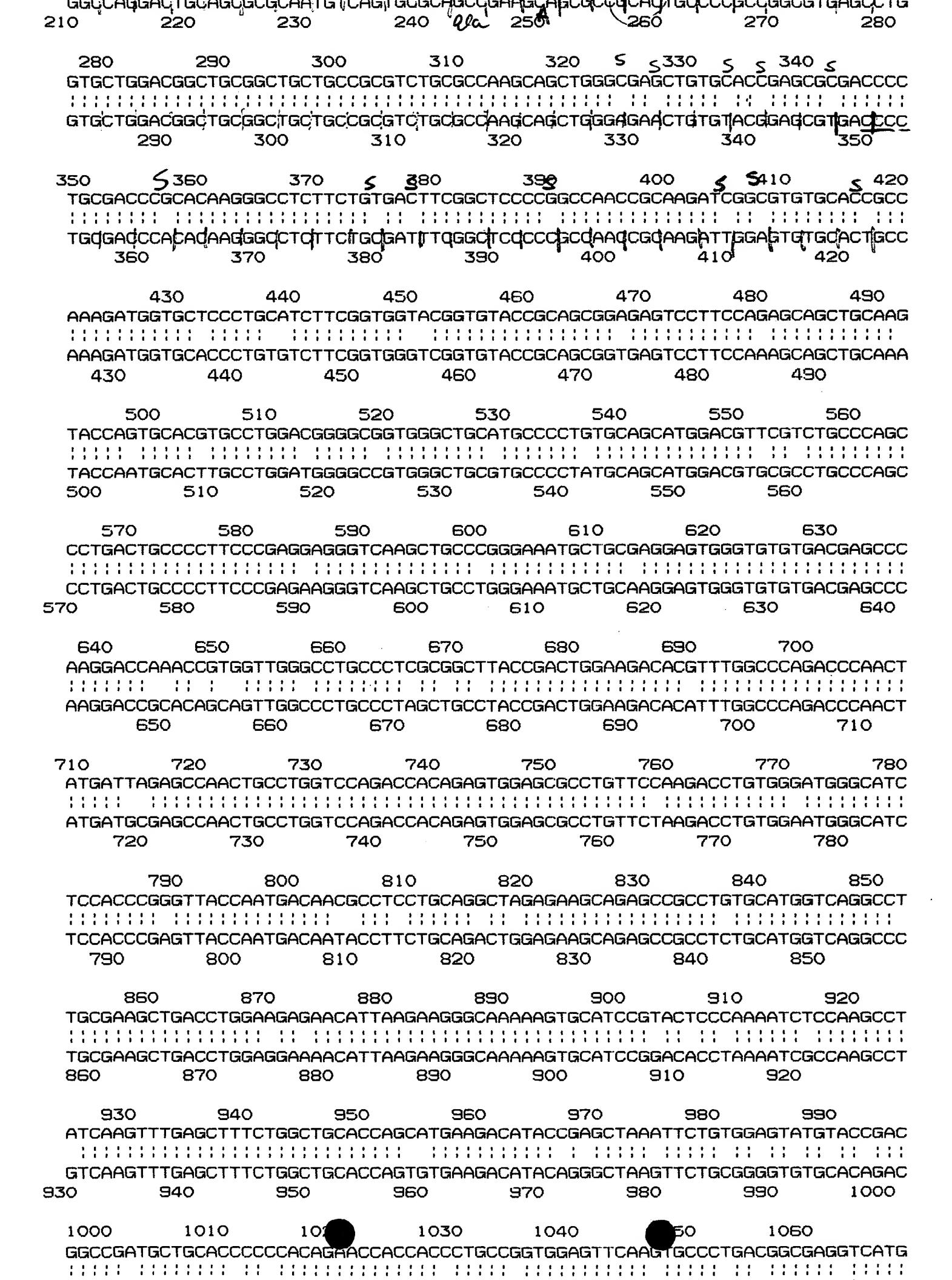
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LOCUS
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                                                          09-JUL-1991
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           M70641
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KEYWORDS
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REFERENCE
              (bases 1 to 4128)
            Ryseck, R. -P., MacDonald-Bravo, H., Mattei, M. -G. and Bravo, R.
  AUTHORS
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            Structure, mapping and expression of fisp-12, a growth factor
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  JOURNAL
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09-JUL-1991
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                                            ROD
LOCUS
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          MUSFISP12B
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DEFINITION
ACCESSION
          M70642
KEYWORDS
          cysteine-rich protein; growth factor-inducible gene.
SOURCE
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  ORGANISM
          Mus musculus
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             (bases 1 to 2267)
REFERENCE
          Ryseck, R. -P., MacDonald-Bravo, H., Mattei, M. -G. and Bravo, R.
  AUTHORS
  TITLE
          Structure, mapping and expression of fisp-12, a growth factor
          inducible gene encoding a secreted cysteine-rich protein
          Cell Growth Differ. 2, 225-233 (1991)
  JOURNAL
          full staff_entry
  STANDARD
                  Location/Qualifiers
FEATURES
    CDS
                  138. . 1184
                  /product="FISP-12 protein"
                  /gene="fisp-12"
                  /codon_start=1
    polyA_signal
                  2224..2230
                 /gene="fisp-12"
                     569 c 582 g
BASE COUNT
                                    546 t
             570 a
ORIGIN chromosome 10
Initial Score
             = 1139 Optimized Score = 1522 Significance =
Residue Identity = 74% Matches
                                    = 1592 Mismatches
                                                         436
             = 100 Conservative Substitutions
Gaps
                                                            0
                              30
                                               50
            10
                    20
                                      40
     CCCGGCCGACAGCCCCGAGACG--ACAGCCCGGCGCGCGTCCCGGTCCCCACCTCCGACCACCGCCAGCGC
       GAATTCCGCCGACAACCCC-AGACGCCACCGCCTGGAGCGTCCAGACACCAACCTCCG-CCCCTGTCCG-AA
                   20
                                                     60
     X 10
                           30
                                   40
                                            50
    70
            80
                    90
                         100
                                      110
                                                  120
                                                        130
   TCCAGGCTCCAGCCGCGCCTCTCGCCCTCTGCACCCTGCTGCATCCTCCTACCGC-GTCCCGATCATG
  70
          80
                  90
                          100
                                110
                                           120
                                                    130
                                                            140
               150 160 170 180
       140
                                                190
                                                         200
   CTCGCCTCCGTCGCAGGTCCCATCAGCCTCGCCT---TGGTGCTCCTCGCCCTCTGCACCCGGCCTGGTACG
                                                     200 Coding
         150
                 160
                         170
                                    180
                                            190
     210
             220
                      230
                             240 S P 250 Rel 2605 S
```

A. Vinelandii nitrogen fixatio

2001

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0.00

2. HATMILOZA

ਠ.	H22KICLI8	Pseudorables virus ichia. 5 ge	2524	100	845	7. 70	0
9.	HUMCANPRA	Human calcium-dependent prote	1154	164	390	7. 46	0
10.	TTHCAAA	T. thermophilus cytochrome caa	1235	164	286	7. 46	0
11.	XELRGEE83	x. laevis rrna external transc	865	162	210	7. 34	0
12.	XELRGMN3	x. laevis 28s rrna non-transcr	1151	162	243	7. 34	0
13.	HS5HCMVCG	Human Cytomegalovirus Strain	229354	158	852	7. 10	0
		**** 6 standard deviations a	above mea	an ****			
14.	HUMRET5	Human mRNA for ret proto-onco	989	154	431	6.86	0
15.	SERERYFGH	S. erythraea 6-deoxyerythromol	2243	152	3 94	6. 74	0
16.	MUSHTF9	Mouse DNA for GC rich region	3725	151	869	6. 69	0
17.	HVBLE	Barley (H. vulgare) lectin mRN	972	150	405	6. 63	0
18.	BLYLEC	Barley root-specific lectin m	972	150	405	6. 63	0
19.	WHTAGGTA	Wheat (T. aestivum) germ agglu	998	149	417	6. 57	0
20.	HUMVIM	Human vimentin gene, complete	1749	148	717	6.51	0

Query sequence being compared: US-07-752-427-1 (1-2075) Number of sequences optimized: 69

Results of the optimized comparison of US-07-752-427-1 (1-2075) with: Data bank : GenBank 69, all entries
Data bank : UEMBL 28_69, all entries

PARAMETERS

Similarity matrix	Unitary	K-tuple	4
Mismatch penalty	. 1	Joining penalty	30
Gap penalty	1.00	Window size	11
Gap size penalty	0. 33		
Cutoff score	124		
Randomization group	0		
Initial scores to save	20	Alignments to save	10
Optimized scores to sav	/e 20	Display context	10

SEARCH STATISTICS

Scores:	Mean 820	Median 821	Standard Deviation 0.00
Times:	CPU 00:00:09.98		Total Elapsed 00:00:16.00
Number of residues:		458541	

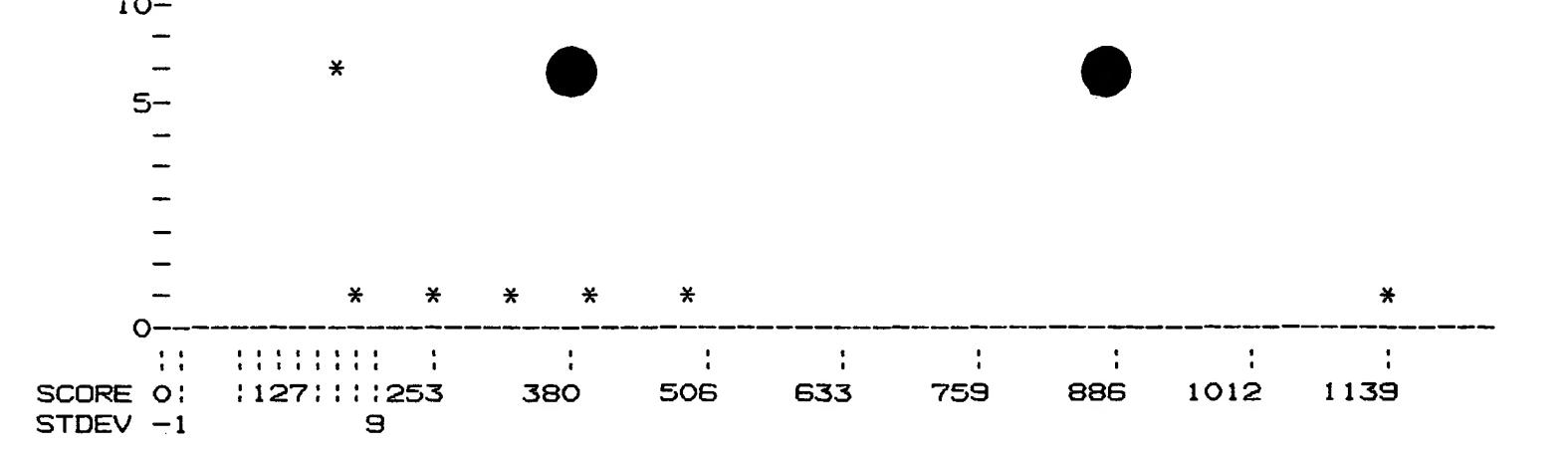
Number of residues: 458541 Number of sequences optimized: 69

The scores below are sorted by optimized score. Significance is calculated based on optimized score.

A 100% identical sequence to the query sequence was not found.

The list of best scores is:

Seque	nce Name	Description	Length	Init. Score	•	Sig.	Frame
1.	MUSFISP12B	Mouse FISP-12 protein (fisp-1	2267	1139	1522	0.00	0
2.	MUSFISP12A	Mouse FISP-12 protein (fisp-1	4128	498	954	0.00	0
3.	HUMFGFAA	Human fibroblast growth facto	3901	146	878	0.00	0
4.	HUMERP	Human erythropoietin gene, co	3398	128	877	0.00	0
5.	MUSHTF9	Mouse DNA for GC rich region	3725	151	869	0.00	0
6.	HUMSRF	Human som response factor (420	134	868	0.00	0
7.	MUSERPA	Mouse erythropoietin gene, co	3891	139	867	0.00	0
8.	CHKCEF	Chicken CEF-10 protein mRNA,	1805	326	865	0.00	0



PARAMETERS

Similarity matrix	Unitary	K-tuple	4
Mismatch penalty	1	Joining penalty	30
Gap penalty	1.00	Window size	11
Gap size penalty	0. 33		
Cutoff score	124		
Randomization group	0		
Initial scores to save	20	Alignments to save	10
Optimized scores to say	ve 20	Display context	10

SEARCH STATISTICS

69

Scores:	Mean	Median	Standard Deviation
	39	38	16.75
Times:	CPU 01:04:13.07		Total Elapsed 01:15:09.00
Number of res	idues:	74447152	
Number of sea	uences searched:	57272	

The scores below are sorted by initial score. Significance is calculated based on initial score.

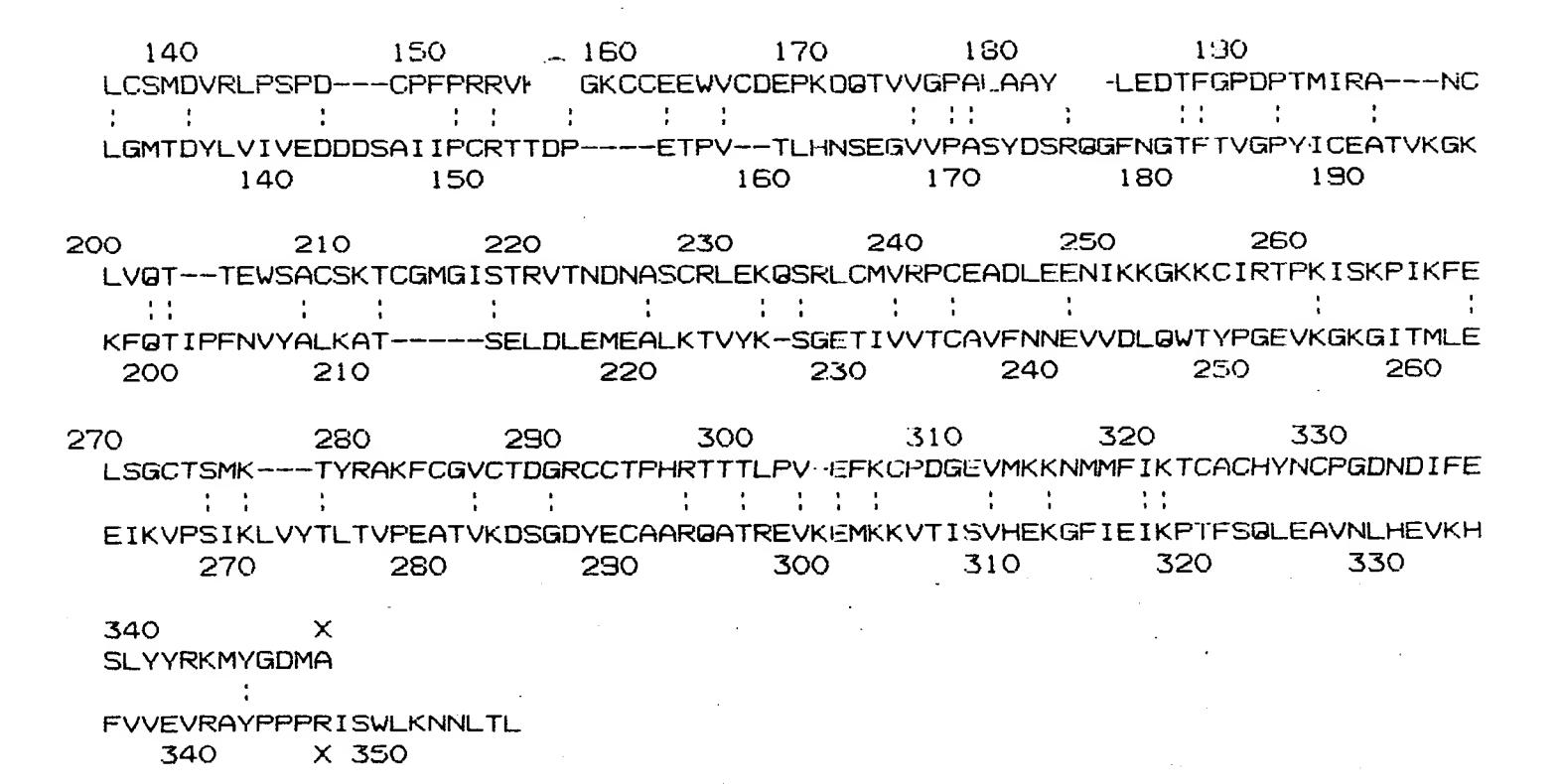
A 100% identical sequence to the query sequence was not found.

The list of best scores is:

Number of scores above cutoff:

Seque	nce Name	Description	Length :	Init. Score	•	Sig. F	rame
		**** 65 standard deviations	above me	an ***	. ×		
1.	MUSFISP12B	Mouse FISP-12 protein (fisp-1	2267	1139	1522	65. 66	0
		**** 27 standard deviations	above me	an ***	* *		
2.	MUSFISP12A	Mouse FISP-12 protein (fisp-1	4128	498	954	27.40	0
		**** 22 standard deviations	above me	an ***	**		
3.	MUSCYR61A	Mouse Cyr61 mRNA, complete co	2018	408	848	°55' 05	0
		**** 17 standard deviations	above me	an ***	: *	•	
4.	CHKCEF	Chicken CEF-10 protein mRNA,	1805	326	865	17. 13	0
		**** 13 standard deviations	above me	an ***	: *		
5.	MMCYR61G	Mouse growth factor inducible	5196	264	833	13. 43	0
		**** 8 standard deviations	above me	an ***	* *		
6.	HUMNFIL6	Human gene for nuclear factor	1910	181	621	8. 48	0
		**** 7 standard deviations	above me	an ***	: X		
7.	PSEPMI	P. aeruginosa pmi gene encodir	1990	170	718	7. 82	0

```
> 0 (
O: :O IntelliGenetics
> 0 (
FastDB - Fast Pairwise Comparison of Sequences
Release 5.4
Results file us-07-752-427-1. res made by maryh on Thu 6 Feb 92 16:25:14-PST.
Query sequence being compared: US-07-752-427-1 (1-2075)
                                               57272
Number of sequences searched:
Number of scores above cutoff:
                                                  69
      Results of the initial comparison of US-07-752-427-1 (1-2075) with:
   Data bank : GenBank 69, all entries
   Data bank : UEMBL 28_69, all entries
100000-
N
U50000-
M
B
Ε
    -**
0
F10000-
S
E 5000-
E
N -
C -
E -
S 1000-
          ¥
   500-
            ¥
   100-
    50-
             ×
               ×
```



```
O: :O IntelliGenetics
> 0 (
FastDB - Fast Pairwise Comparison of Sequences
Release 5.4
Results file us-07-752-427-2-a-geneseq. res made by maryh on Thu 6 Feb 92 15:28:31-f
Query sequence being compared: US-07-752-427-2 (1-349)
Number of sequences searched:
                                           14140
Number of scores above cutoff:
                                            3377
     Results of the initial comparison of US-07-752-427-2 (1-349) with:
  Data bank : A-GeneSeq 5, all entries
 10000-
N
U 5000-
M
8
Ε
                 X
                              ×
0
F 1000-
                                  X
  500-
                                          ×
                                              ×
   100-
         *
                                                  ¥
   50-
    10-
                                                           ¥
```

1 1	1 1	•	1 1		1		1 1	1	1
SCORE 0:	2 :	3	51	7:	8	: 10	:12	13	15
STDEV -1	0			2		3	4	5	

PARAMETERS

Similarity matrix Ur	nitary	K-tuple	2
Mismatch penalty	1	Joining penalty	20
Gap penalty	1.00	Window size	11
Gap size penalty	0.05		
Cutoff score	6		
Randomization group	0		
Initial scores to save	20	Alignments to save	10
Optimized scores to save	20	Display context	10

SEARCH STATISTICS

Scores:	Mean 4	Median 5	Standard Deviation 2.04
Times:	CPU 00:00:42.09		Total Elapsed 00:01:27.00
Number of residues		2168208	
Number of sequence		14140	
Number of scores a	above cutoff:	3377	

The scores below are sorted by initial score. Significance is calculated based on initial score.

A 100% identical sequence to the query sequence was not found.

The list of best scores is:

				Init.	Opt.		
Seque	nce Name	Description	Length	Score	Score	Sig.	Frame
		**** 5 standard deviations	above me	an **;	+ *		
1.	P93284	Sequence of clone HIV-2 SBL/I	3025	15	57	5. 39	0
		**** 4 standard deviations	above me	an **	{ X		
2.	P81243	Human spleen trypsin III (try	247	14	37	4. 90	0
3.	R11741	Granulocyte colony stimulatin	783	14	32	4. 90	0
4.	R11742	Clone 25-1 encoded human G-CS	801	14	35	4. 90	0
5.	R08039	Cathepsin D fragment of fibro	993	14	45	4. 90	0
6.	P70373	Human fibronectin gene produc	2327	14	45	4. 90	0
7.	R12185	Protease inhibitor with varia	114	13	17	4.41	0
8.	P94681	Amino acid sequence encoded b	3 48	13	25	4. 41	0
		**** 3 standard deviations	above me	an ***	(
9.	R10109	Trigramin-beta 1	72	12	20	3. 92	0
10.	P91320	New trigramin peptide isolate	72	12	20	3. 92	0
11.	R10110	Trigramin-beta 2.	73	12	20	3. 92	0
12.	R10106	Trigramin-gamma.	73	12	19	3. 92	0
13.	P96395	Albolabrin.	73	12	19	3. 92	0
14.	R06395	Albolabrin.	73	12	19	3. 92	0
15.	R07329	Amb a I/Antigen E encoded by	92	12	20	3. 92	0
16.	R07330	Amb a I/Antigen E encoded by	94	12	21	3. 92	0
17.	R07455	N-terminal sequence of anti-p	120	12	22	3. 92	0
18.	P91901	Deduced sequence of porcine e	203	12	34	3. 92	0
19.	P91376	Porcine endothelin III.	203	12	34	3. 92	0
20.	P90502	Gp. B Eimeria tenella immunoge	237	12	36	3. 92	0
		_					

Results of the optimized comparison of US-07-752-427-2 (1-349) with: Data bank : A-GeneSeq 5, all entries 1000-Ν U 500-M В E R 0 ¥ F 100- * × ¥ × ¥· × X S × * * X × × X Ε * * 50-0 × U X Ε × NC E S 10-¥ * * × × STDEV-3 39 42 SCORE36 51 54 57 E:O 62 45 48 PARAMETERS 2 Similarity matrix K-tuple Unitary Mismatch penalty Joining penalty 20 Window size Gap penalty 11 1.00 Gap size penalty 0.05 Cutoff score 6 Randomization group 0 Initial scores to save Alignments to save 10 20 Display context 10 Optimized scores to save 20 SEARCH STATISTICS Standard Deviation Scores: Mean Mediari 49 45 3.08 Times: CPU Total Elapsed 00:01:28.89 00:02:56.00 Number of residues: 1361708 Number of sequences optimized: 3377

The scores below are sorted by optimized score.

3311

Number of sequences uptimized:

-significance is calculated based on optimized score.

A 100% identical sequence to the query sequence was not found.

The list of best scores is:

				Init.	Opt.		
Sequer	nce Name	Description	Length	Score	Score	Sig.	Frame
		**** 5 standard deviations	above me	ean ***	*		
1.	P92069	Human muscarinic acetylcholin **** 4 standard deviations		10 ean ***	63 *	5. 84	0
2.	P60053	Sequence of von Willebrand fa	a 2813	11	. 59	4. 54	0
3.	P60462	Sequence of human von Willeb	r 2813	7	59	4. 54	0
		**** 3 standard deviations	above me	ean ***	*		
4.	P93284	Sequence of clone HIV-2 SBL/	I 3025	15	57	3.89	0
5.	P80674	NtrA gene product.	523	6	57	3. 89	0
6.	R10534	Human 160kD mediator of infla	a 1427	10	57	3, 89	0
7.	P93357	Sequence of the catalytic dom	m 1522	9	57	3. 89	0
8.	R12108	N-terminal deleted adenyl cyc	1445	9	56	3. 57	0
9.	R08267	Platelet derived growth facto	1089	6	56	3. 57	0
10.	R06910	Alpha type PDGF receptor ded	u 1089	6	56	3. 57	0
11.	P94365	Sequence of part of adenylate	e 1705	9	56	3. 57	0
12.	R07683	gag protein precursor, p75gag	g 636	6	56	3. 57	0
13.	P80810	Sequence of pol protein of H	1014	6	55	3. 25	0
14.	P90599	Human retinoblastoma.	970	6	55	3. 25	0
15.	R07713	Human low density lipoprotein	900 n	8	55	3. 25	0
16.	P60057	Factor IX/Factor VII fusion	453	9	55	3. 25	0
17.	P90180	Tissue plasminogen activator	534	6	55	3. 25	0
18.	P60056	Factor VII peptide encoded by	y 466	9	55	3. 25	0
19.	R08031	Adenyl cyclase from Bordetel	1706	9	55	3. 25	0
20.	P60055	Partial Factor VII peptide.	371	9	55	3. 25	0

1. US-07-752-427-2 (1-349)

P92069 Human muscarinic acetylcholine m2 receptor gene.

- P92069 standard; protein; 2135 BP. ID
- AC P92069;
- DT 27-SEP-1989 (first entry)
- DE Human muscarinic acetylcholine m2 receptor gene.
- KW Muscarinic acetylcholine receptor; drug screening; probes; m2; ss.
- os Homo sapiens.
- FH Location/Qualifiers Key
- 120. . 1517 FT CDS
- /*tag= a FT
- FT /product=MAR subtype m2
- FT misc_signal 74
- FT /*tag= b
- FT /label=splice acceptor site
- FT /note=defines the 5' end of the exon
- FT 74..2096 polyA_site
- FT /*tag= c
- PN US7241971-A.
- PD 14-MAR-1989.
- PF 08-SEP-1988; 241971.
- PR 08-SEP-1988; US-241971.
- PA (USSH) Nat Inst of Health.
- DR WPI; 89-165452/22.
- DR P-PSDB; P96203.
- Cloned genes for muscarinic acetylcholine receptors -PT
- PT for drug screening and diagnostic use.
- PS Disclosure; p; English.
- CC The sequence may be useful for synthesis of hybridisation probes for
- CC diagnostic use. The genes are cloned by screening a rat cerebral cortex

```
CC
      strand of porcine brain (m1) cDNA (modified at positions 5, 38 and 53);
 CC
      identifying cDNA clones by hybridisation with BamHI or XhoI digests of
 CC
      plasmid DNA from 12 cultures contg. up to 500000 independent clones;
 CC
      reculturing until a pool contg. less than 5000 indep. clones with a
      single hybridisation band are identified, and isolating individual clones
 CC
 CC
      by colony hybridisation.
 CC
      Stable cell lines are produced by transfecting Chinese hamster ovary cells
      (CHD-K1) with various pCD vectors contg. the gene inserts.
 CÇ
 CC
      See also N92068-N92067 and N90086.
 SQ
      Sequence
                 2135 BP;
                                      462 C;
                                               448 G;
                            629 A;
                                                         596 T;
Initial Score
                            Optimized Score
                                                   63 Significance =
                        10
                                                        Mismatches
Residue Identity =
                      20%
                           Matches
                                                    75
                                                                         266
                            Conservative Substitutions
Gaps
                       20
                                                                           0
                                                                50
                                 20
                                           30
                                                      40
                                                                          60
                       10
              X
              MTAASMGPVRVAFVVLLALCSRPAVG@NCSGPCRCPDEPAPRCPAGVSLVLDGCGCCRVCAK
    AAAGTCAACCGCCACCTCCAGACCGTCAACAATTACTTTTTATTCAGCTTGGCCTG-TGCTGACCTTATCAT
                                       300
                                                 310
        270
                             290
                                                             320
                                                                       330
              X
                  280
                                                     110
                                                               120
                                                                            130
          70
                    80
                               90
                                          100
    QLGELCTERDPCDPHKGLFCDFGSPANRKIGVCT--AKDGAPCIFGGTVYRSGESFQSSCKYQCTCLDG--A
                                     1 1
                                                   1 1 1
    AGGTGTTTTCTC-CATGAACTTG-TACACCCTCTACACTGTGATTGGTTACTGGCCTTTGGGACCTGTGGTG
                                        370
                                                  380
                                                             390
                                                                       400
       340
                  350
                              360
                                          170
                                                                           200
           140
                     150
                                160
                                                       180
                                                                 190
    VGCMPLCSMDVRLPSPDCPFPRRVKLPGKCCEEWVCDEPKDQTVVG---PALAAYRLEDTFGPDPTMIRANCL
    TGTGACCTTTGGCTAGCCCTGGACTATGTGGTCAGCAATGCCTCAGTTATGAATCTGCTCATCAT--CAGC-
       410
                 420
                           430
                                      440
                                                450
                                                           460
                                                                       470
                                                                           270
           210
                     220
                                230
                                                     250
                                                                 260
                                          240
    VØTTEWSACSKTCGMGISTRVTNDNASCRLEKØSRLCMVRPCEADLEENIKKG--KKCIRTPKISKPIKFEL
    -TTTGACAGGTACTTCTGT-GTCACAAAACCTCTGACCTACCCAGTCAAGCGGACCACAAAAATGGCAGGTA
                                                   520
                                         510
         480
                   490
                               500
                                                              530
                                                                        540
                                                     320
                                           310
                                                                   330
           280
                      290
                                 300
    SGCTSMKTYRAKFCGVCT-DGRCCTPHRTTTLPVEFKCPDGEVMKKNMMFIKTC---ACHYNCPGDNDIFES
                  1 1 1 1 1 1 1 1 1 1
                                 1 1 1
    TGATGATTGCAGCTGCCTGGGTCCTCTCTTTCATCCTCTGGGCTCCAGCCATTCTCTTCTGGCAGTTCATTG
        550
                  560
                            570
                                       580
                                                 590
                                                            600
                                                                      610
   340
    LYYRKMYGDMA
    TAGGGGTGAGAACTGTGGAGG
      620
              X 630
2. US-07-752-427-2 (1-349)
                Sequence of von Willebrand factor (VWF).
   P60053
 ID
      P60053 standard; Protein; 2813 AA.
 AC
     P60053;
DT
      22-JUL-1991 (first entry)
DE
      Sequence of von Willebrand factor (VWF).
KW
      Vascular injury; platelet plug formation.
     Homo sapiens.
OS.
 PN
     EP-197592-A.
PD
     15-0CT-1986.
PF
      26-MAR-1986; 200518.
```

CUNH IIDITATY WITH a probe based on nucleurides I/O-223 of the non-coaring

PR

01-APR-1985; NL-000961.

```
(FRIN-) STICHT FRINDER RANT.
PA
PI
     Pannekoek H, Verwey (
                             Diergaarde PJ, Hart MHL;
     WPI; 86-273504/42.
DR
DR
     N-PSDB; N60061.
     Recombinant cDNA plasmid or phage - contg. C-DNA fragment which
PT
     codes for biological activity of human von Willebrand factor
PT
     Disclosure; Fig 3; 37pp; English.
PS 
CC
     vWF (glyco) protein having the AA sequence corresponding to the
     nucleotide sequence of 2518-8667 or 295-2517 of N60061 is claimed.
CC
     Also claimed are new microorganisms, animal cell or human cell contg.
CC
      the recombinant cDNA plasmid or phage; e.g. strain E. coli DH1 contg.
CC
      the recombinant cDNA plasmid pSP8800vWF is deposited as C.B.S. No
CC
 CC
      163.86.
SØ
     Sequence
                2813 AA;
      160A; 137R; 101N; 160D; 0 B; 217C; 142Q; 181E; 0 Z; 207G; 77 H;
SØ
     97 I; 233L; 104K; 55 M; 93 F; 173P; 201S; 146T; 27 W; 81 Y; 221V;
50
                                                 59 Significance =
                          Optimized Score
                      1 1
Initial Score
                                                     Mismatches
Residue Identity =
                     20%
                          Matches
                                                                      248
                                                 77
                          Conservative Substitutions
                      43
                                                                        0
Gaps
                                                                50
                                                      40
                      10
                                 20
                                           30
              X
             MTAASMGPVRV-AFVVLLALCSRPAVGQ-NCSGPC-RCPDEPAPRCPAGVSLVLDGCGCCRV
   NYNGN@GDDFLTPSGLAEPRVEDFGNAWKLHGDC@DL@K@HSDPCALNPRMTRFSEEACAVLTSPTFEACHR
                                             570
                                                       580
                                                                 590
             X 540
                                   560
     530
                         550
                                                     110
                                                                         130
                                 90
                                                               120
   60
                                           100
             70
                       80
    CAKBLGELCTER-DPCDPHKGLFCDFGSPANRKIGVCTAKDGAPCIFGGTVYRSGESFØSSCKYØCTCLDGA
    AVSPLPYLRNCRYDVCSCSDGRECLCGALA----SYAAACAGRGVRVAWREPGRCELNCPKG@-VYL@CG
                                                 640
                                                           650
                                                                      660
             610
                                       630
    600
                       620
                                                                       190
                                         160
                                                   170
                                                             180
                               150
           140
                          ----CPFPRRVKLPGKCCEEWVCDEPKD@TVVGPALAAYRLEDTFGPDP
    VGCMPLCSMDVRLPSPD---
    TPCNLTC---RSLSYPDEECNEACLEGCFCP----PGLYMDERGDCVPKA@CPCYYDGEIF@PEDIFSDHH
                                                                  720
                               690
                                              700
                                                        710
                     680
            670
                                                              250
                                      230
                                                    240
                  210
                            220
         200
    TMIRANCLV@TTEWSACSKTCGMGISTRVTNDNASCRLEK@S---RLCMV-RPCEADLEENIKKGKKCIRTP
                                                TM---CYCEDGFMHCTMSGVPGSLLPDAVLSSPLSHR-SKRSLSCRPPMVKLVCPAD---NLRAEGLECTKT-
                                                770
                                                             780
                            750
                                      760
        730
                  740
                                                                       330
                                        300
                                                  310
                                                            320
                    280
                              290
           270
    KISKPIKFELSGCTSMKTYRAKFCGVCTDGRCCTPHRTTTLPVEFKCPDGEVMKKNMMFIKTCACHYNC-PG
                             1 1 1
    CONYDLECMSMGCVS--GCLCPPGMVRHENRCVALERCPCFHQGKEYAPGETVKIGCNTCVCRDRKWNCTDH
                               820
                                                             850
                                                                       860
                     810
                                         830
                                                   840
         800
          340
   DNDIFESLYYRKMYGDMA
    VCDATCSTIGMAHYLTFDGLKYLFPGEC
         870
                   880
3. US-07-752-427-2 (1-349)
               Sequence of human von Willebrand Factor (VWF) prec
   P60462
```

FH

(AKIE-) SIICHI AKIEND FUNDS.

DT 25-JUN-1991 (first entry)
DE Sequence of human von Willebrand Factor (VWF) precursor.

P60462 standard; Protein; 2813 AA.

ID

AC

P60462;

```
05
     Homo sapiens.
PN
     WD8606096-A.
PD
     23-0CT-1986.
PF
     10-APR-1986; U00760.
PR
     11-APR-1985; US-722108.
PA
     (CHIL-) CHILDRENS MED CENT.
PA
     (GINS/) GINSBURG D.
ΡI
     Ginsburg D. Orkin SH. Kaufman RJ;
DR
     WPI; 86-291663/44.
DR
     N-PSDB; N60404.
PT
     Pure Von Willebrand Factor - produced using an expression vector
     including a DNA sequence encoding functional VWF protein
PT
     Disclosure; Table 2, Pages 18-36A; 54pp; English.
PS
CC
     cDNA clones pVWH33, pVWH5 and PVWE6 which span 9 kb pairs of DNA and
CC
     encompass the entire protein coding region of VWF, were selected to
     construct full length cDNA (N60404). The pure VWF produced is useful
CC
CC
     in the treatment of von Willebrand's disease (VWD) and the patients
     with chronic renal failure whose abnormal bleeding times are
CC
CC
     corrected by crude cryoprecipitate. Pure VWF can also be used to
CC
     carry, stabilise and improve the therapeutic efficacy of factor
CC
     VIII:C.
 20
               2813 AA;
     Sequence
SØ
     154A; 143R; 98 N; 155D; O B; 234C; 133Q; 181E; O Z; 205G; 70 H;
SQ
     95 I; 227L; 108K; 56 M; 89 F; 176P; 207S; 151T; 26 W; 79 Y; 226V;
Initial Score
                        Optimized Score =
                                             59
                                                 Significance =
                                                               4. 54
                                                 Mismatches
                                                                259
                        Matches
Residue Identity =
                                             76
                    20%
                                                                  0
Gaps
                    28
                        Conservative Substitutions
            X
                                        30
                                                 40
                                                          50
                                                                   60
                     10
                              20
            MTAASMG-PVRVAFVVLLALCSRPAV-G@NCSGPCRCPDEPAPRCPAGVSLVLDGCGCCRVC
                                  1 1 1
   CLPDKVCVHRSTIYPVG@FWEEGCDVCTCTDMEDAVMGLRVA@CS@KPCE--DSCRSGFTYVLHEGECCGRC
            X 2440
     2430
                                2460
                                         2470
                                                    2480
                                                              2490
                       2450
             70
                      80
                                90
                                         100
                                                   110
                                                            120
   AK@LGELCT---ERDPCDPHKGLFCDFGSPAN-RKIGVCTAKDGAFCIFGGTV-YRSGESF@SSCKY@CTCL
   LPSACEVVTGSPRGDSQSSWKSVGSQWASPENPCLINECVRVKEEVFIQQRNVSCPQLEVPVCPSGFQLSCK
              2510 2520 2530 2540
                                                   2550
     2500
   130 140 150 160 170 180
   DGAVGCMPLCSMDVRLPSPDCPFPRRVKLPGKCCEEWVCDEPKD@TVVGPALAAYRLE---DTFGPDPTMIR
                      TSA--CCPSC---RCERMEACMLNGTVIGPGKTVMIDVCTTCRCMV@VG-VISGFKLECRKTTCNPCP--LG
                                   2600 2610 2620
                 2580 2590
     2570
    200 210 220 230 240 250
   ANCLV@TTEWSACSKTCGMGISTRVTNDNASCRLEK@SRLCMVRPCEADLEENIKKGKKCIRTPKISKPIKF
   YKEENNTGECCGRCLPTACTIQLRGGQIMTLKRDETLQDGCDTHFC--KVNERGEYFWEKRVTGCPPFDEHKC
                           2660 2670 2680
         2640
                  2650
                                                       2690
  270 280 290 300 310 320 330
   ELSGCTSMKTYRAKFCGVCTDGRCCTPHRTTTL-PVEFKCPDGEVMKKNMMFIKTCACHYNCPGD-ND--IF
   LAEGGKIMK-IPGTCCDTCEEPEC--NDITARL@YVKVGSCKSEVEVDIHYC@GKCASKAMYSIDINDV@D@
                             2730 2740
        2710
                  2720
                                               2750 2760
    340
   ESLYYRKMYGDMA
   CSCCSPTRTEPM@VALHCTNGSV
                  2790
         2780 X
```

Chronic renal fallure; therapy; factor vill C.

KW.

```
Sequence of lone HIV-2 SBL/ISY.
  P93284
     P93284 standard; protein; 3025 AA.
 ID
 AC
     P93284;
 DT
     O6-APR-1990 (first entry)
 DE
     Sequence of clone HIV-2 SBL/ISY.
 KW
     HIV-2; proviral clone HIV-2 SBL/ISY;
 05
     Human immunodeficiency virus 2.
 PN
     US7331212-A.
 PD
     29-AUG-1989.
 PF
     31-MAR-1989; 331212.
 PR
     31-MAR-1989; US-331212.
 PA
     (USSH) US Dept. Health and Human Services.
     Franchini G, Wong-Staal F, Gallo R;
 PI
 DR
     WPI; 89-339698/46.
 DR
     N-PSDB; N92119.
 PT
     Complete human immunodeficiency type 2 proviral clone - used to generate
     animal model for function studies of HIV genes in vivo.
PT
     Disclosure; Fig. 5; 43pp; English.
 PS
     The protein is encoded by the second reading frame of HIV-2 SBL/ISY, a
 CC
 CC
     proviral clone of HIV-2.
 50
     Sequence
               3025 AA;
            278 R; 126 N;
 26
     143 A;
                           104 D; O B; 96 C; 191 Q; 115 E; O Z;
                                                                 244 G;
     95 H; 153 I; 224 L; 246 K; 65 M;
SB
                                      63 F;
                                             160 P;
                                                    291 S; 193 T; 53 W;
S@
     80 Y; 105 V;
Initial Score
                        Optimized Score =
                                                Significance = 3.89
                  15
                                             57
Residue Identity =
                        Matches
                   20%
                                                Mismatches
                                                                256
                                             75
                                                            ----
Gaps
                    41
                        Conservative Substitutions
                                                                 0
            X
                   10
                            20
                                     30
                                              40
                                                             50
            MTAASMGPVRVAFVVLLALCSRPAVG@NCSGPCRCPDEPA----PRCPAGVSLVLDGCGC
   MDSHPETSGCPKLGGTNLPRNKDQTLMA---NRKDDTHGRSTVDRISRSRAGGKQNYLKPGTGGTLLPRGKRV
    980
             990
                     1000
                                1010
                                         1020
                                                           1040
                                                  1030
                                  90 100 110
                70 80
   CRVCAK---QLGELCTERDPCDPHKGLFCDFGSPANRKIGVCTAKDGAPCIFGGTVYRSGESFØSSCKYØCT
   RSNSSKGSRØSVDISTPGRKNSKSRKICKDKKYPYØRGØVVGT---GSSENRKRSTSHLGTNTKISPTSRKR
         1060 1070 1080
   1050
                                    1090 1100 1110
     130 140 150 160 170 180
                                                                190
   CLDGAVGCMPLCS-MDVRLPSPDCPFPRRVKLPGKCCEEWVCDEPKD@TVVGPALAA-----YRLEDTFGPD
   DLGTVVG--LLASDMDPRLGLRIHPTVG@VSIPGKRSYTRRRD-----LLHGWIL@AIKRGKSRIYNRRKR@
            1130 1140 1150
    1120
                                          1160 1170
                 210 220
        200
                                   230
                                                240
                                                         250
   PTMIRANCLV@TTEWSACSKTCGMGISTRVTNDNASCRL----EK@SRLCMVRPCEADLEENIKKG-KKCIR
   SKDIRAN--Y@PT--SRIRSLCNGSNRLRSKS@YCS-RLTVCNGNSNRPTGIRENSKNYRRDDKKGSNLCCM
                              1210 1220
          1190
                   1200
                                              1230
  260
                      280 290 300
            270
                                                  310
   TP-KISKPIKFELSGCTS-MKTYRAKFCGVCTDGR-CCTPHRTTTLPVEFKCPDGEVMKKNMMFIKTCACHY
   GPGPGRHRRKSRNPLSKSGHØTSIIPRENRARSGRTWKISØCKRTSPVWITØPGGKTNSKHMCPVPTKRGSY
  1250 1260
                    1270 1280
                                    1290
                                             1300
   330 340
   NCPGDNDIFESLYYRKMYGDMA
```

4. US-07-752-427-2 (1-349)

-TWASKCRTRHLANGLHTLRMKNHYSSSTCCK

```
1320
           1330
                    1340
                              1350
5. US-07-752-427-2 (1-349)
               NtrA gene product.
  P80674
 ID
     P80674 standard; protein; 523 AA.
 AC
     P80674;
 DT
     24-0CT-1990 (first entry)
 DE
     NtrA gene product.
 KW
     NtrA; sigma factor; transcriptional activation; nitrogen assimilation;
     fixation; nif operon; C4-dicarboxylate transport.
 KW
     Rhizobium meliloti.
 05
 PN
     EP-292984-A.
 PD
     30-NOV-1988.
 PF
     27-MAY-1988; 108482.
 PR
     29-MAY-1987; US-055228.
 PA
     (GEHO-) Gen Hospital Corp.
 PI
     Ronson C, Ausubel F;
 DR
     WPI; 88-339561/48.
 DR
     N-PSDB; N81266.
 PT
     Cloned rhizobium meliloti ntrA gene - for use in C4-dicarboxylate
     transport, nitrate assimilation, symbiotic nitrogen fixation and
PT
PT
     identifying pathogenic genes.
PS
     Claim 1; Page 12; 19pp; English.
 CC
     The ntrA gene product is a transcriptional activator. It controls the
CC
     processes of nitrate assimilation, symbiotic nitrogen fixation and
CC
     C4-dicarboxylate transport, working in conjunction with NtrC, NifA and
     DCtD resp. It can also be used to identify genes involved in
CC
     pathogenesis.
CC
     See also P82126.
CC
20
     Sequence
               523 AA;
50
     60 A; 43 R; 15 N;
                         43 D;
                               OB; 4C; 250;
                                                 39 E; O Z; 29 G;
                                                                    12 H;
     23 I; 58 L; 15 K; 13 M;
                               17 F; 25 P; 37 S; 21 T; 3 W; 8 Y; 33 V;
SØ
Initial Score
                         Optimized Score =
                                              57
                                                  Significance =
                      6
                                                                 3. 89
Residue Identity =
                         Matches
                                              75
                                                  Mismatches
                                                                  257
                    20%
                                                              ----
                         Conservative Substitutions
                     39
Gaps
                                        30
                             20
             X
                   10
             MTAASMGPVRVAFVVLLALCSRPA--VGQNCSGPCRCPDEP--APRCPAGVSLVLDGCGCCR
   LHLRQSQSLVMTPQLMQSIQLLQMNHLELSHFIAQEVEKNPLLEVQPADEPTISDREDAGPHPAETG-GETD
                                 40
                                           50
     10
                        30
                                 90
   60
              70
                        80
                                         100 110
   VCAK@--LGELCTERDPCDPHKGLFCDFGSPANRKIGVCTAKDGAPCIFGGTVYRSGESF@SSCKY@CTCLD
   EAAGGSDLYDSAMSRSGERLSEGLDADF---AN-VFPDDTAPQRADAPELLGQVKSMPGAGDAEGY--DLDD
                      100
    80
                                   110
                                                    130
             90
                                            120
                    150 160 170 180 190
  130 140
   GAVGCMPL-CSMDVRLP----SPDCPFPRRVKLPGKCCEEWVCDEFKD@TVVGPA--LAAYRLEDTFGPDP
   FVGGRKTLRETLAEGLPFALSAVSDRLIARYFID@LDDAGYLHADLAETAETLGAAGEDVARVLHVL@QFDP
                         170
       150
                                          190
                160
                                180
                                                      200
                     210 220 230 240 250
        200
   TMIRANCLV@TTEVSA----CSKTCGMGISTRVTNDNASCRLEK@SRLCMVRPCEADLEENIKKGKKCIRTP
   PGVFARTL---GECLAI@LRARNRLDPAMEALVANLELLARRDFASLKKICGVDEEDLIDMLAEIRKLDPKP
                 230
                                    250
     220
                          240
                                             260
                                                       270
                                                           320
           270
                    280
                             290 300
   KIS-KPIKFELSGCTSMKTYRAKFCGVCTDGRCCTPHRTTTLPVEFKCPD-GEVMKKNMMFIKTCACHYNC
```

```
RIZE INVEE - HITTOUVVAH----HUDRING MENEUM DHENKALANDAH I FIZK ZZK NZREMHENEC
       290
                                   310
                   300
                                             320
                                                                 340
                                                       330
   330
             340
                        X
    -PGDNDIFESLYYRKMYGDMA
             :: :
    L@NANWLTRSLD@RARTIMKVASEIVR@@DA
  350
            360
                      370
                                 380
6. US-07-752-427-2 (1-349)
   R10534
                Human 160kD mediator of inflammation protein.
 ID
      R10534 standard; Protein; 1427 AA.
 AC
      R10534;
 DT
                   (first entry)
      12-APR-1991
 DE
      Human 160kD mediator of inflammation protein.
 KW
      Mediator of Imflammation; cytokine; Hodgkin's lymphoma; MRP-160.
 05
      Homo sapiens.
 PN
      EP-412050-A.
 PD
      06-FEB-1991.
 PF
      26-JUN-1990; 810481.
 PR
      05-JUL-1989; GB-015414.
 PA
      (CIBA ) CIBA GEIGY AG.
      Odink KG, Tarcsay L, Bruggen J, Wiesendanger W, Cerletti N;
 PI
 PI
      Sorg C, Dewolf-Peeters C, Delabie J;
      WPI; 91-038913/06.
 DR
 DR
      N-PSDB; @10378.
      160 kD human polypeptide mediator or precursor of inflammation -
 PT
 PT
      polyclonal or monoclonal antibodies to polypeptide treat and
 PT
      diagnose chronic inflammation and hodgkins lymphoma
 PS
      Claim 3; Page 32; 47pp; English.
      The protein is a cytokine used to treat chronic inflammatory
 CC
 CC
      conditions. It is prepared by chromatographically purifying an
 CC
      optionally pre-purified cell extract, cell supernatant or cell
      filtrate of stimulated normal human leucocytes or human embryonic
 CC
      epithelial lung cells. Alternatively, the protein can be produced
 CC
 CC
      by microorganisms or continuous mammalian cell lines, transformed
 CC
      with plasmids encoding MRP-160. The invention also covers the
 CC
      polypeptide fragment from amino acids 878-1427 and derivatives of
 CC
      the protein in which the amino and/or hydroxyl functions are
 CC
      glycosylated or acylated and have mol. wt.s of 190 and 140kD,
 CC
      respectively.
 SØ
      Sequence
                 1427 AA;
      115A; 67 R; 59 N; 70 D; 0 B; 14 C; 90 Q; 199E; 0 Z; 49 G; 26 H;
 SØ
 SB
      45 I; 156L; 153K; 33 M; 29 F; 34 P; 121S; 88 T; 5 W; 9 Y; 65 V;
Initial Score
                       10 Optimized Score
                                                   57 Significance =
                                                                       3.89
Residue Identity =
                      19% Matches
                                                       Mismatches
                                                                        258
                                                   73
Gaps
                           Conservative Substitutions
                       39
                                                                          0
              X
                      10
                                20
                                                                 50
                                          30
                                                       40
                                                                           60
              MTAASMGPVRVAFVVLLALCSRPAVG@NCSGPCRCPDE--PAPRCPAGVSLVLDGCGCCRVC
    SNLTKTASESISNLSEAGSIKKGERELKIGDRVLVGGTKAGVVRFLGETDFAKGEWCGVELDEPLGKNDGAV
      190
              X 200
                                    220
                          210
                                              230
                                                         240
                                                                   250
              70
                        80
                                    90
                                              100
                                                        110
                                                                   120
   AKQLGELC--TERDPCDPHKGLFCDFGS--PANRKIGVCTAKDGAFCIFGGTVYRS-GESFQSSCKYQCTCL
   AGTRYF@C@PKYGLFAPVHKVTKIGFPSTTPAKAK---ANAVRRVMATTSASLKRSPSASSLSSMSSVASCV
   260
              270
                        280
                                  290
                                               300
                                                          310
                                                                    320
    130
                  140
                            150
                                      160
                                                170
                                                           180
                                                                     190
   DGAVGCMPL----CSMDVRLPSPDCPFPRRVKLPGKCCEEWVCDEPKD@TVVGPALAAYRLEDTFGPDPTMI
```

```
22Kh2KIPITIE 122KAUKKI2PILUFERFUKEKRMHIFMFUTEKUFEKUKU-12HAPITERFTUKUK
                         350
     330
               340
                                   360
                                             370
                                                        380
                                                                   390
                                     230
                                                240
                                                                         260
                210
                          220
                                                               250
      200
    RANCLV@TTEWSACSKTCGMGISTRV-TNDNASCRLEK@SRLCMVRPCEADL----EENIKKGKKCIRTPK
    GHD@HVLELE-----AKMD@LRTMVEAADREKVELLN@-LEEEKRKVE-DL@FRVEEESITKGDLET@TKL
    400
                              420
                                        430
                                                     440
                                                               450
                                                                         460
                    410
                    280
                              290
                                                                320
                                                                          330
                                           300
                                                      310
          270
    ISKPIKFELSGCTSMKTYRAKFCGVCTDGRCCT---PHRTTTLPVEFKCPDGEVMKKNMMFIKTCACHYNCP
                                                         1 1
                   | |
    EHARIKELEGSLLFEKTKADKLGRELEDTRVATVSEKSRIMELEKDLALRVGEV-----AELRRRLESNKP
                                                   510
          470
                    480
                                                                   520
                              490
                                        500
            340
                       X
    -GDNDIFESLYYRKMYGDMA
     1 1 1 1 1 1 1
    AGDVDMSLSLL@EISSL@EKLEVTRTDH@R
    530
              540
                        550
7. US-07-752-427-2 (1-349)
                Sequence of the catalytic domain of Bordetella per
   P93357
      P93357 standard; protein; 1522 AA.
 ID
      P93357;
 AC
      22-MAR-1991 (first entry)
 DT
      Sequence of the catalytic domain of Bordetella pertussis
 DE
      adenylate cyclase (AC).
 DE
      Vaccine; ss.
KW
05
      Bordetella pertussis.
      FR2618453-A.
 PN
      27-JAN-1989.
 PD
PF
      24-JUL-1987.
 PR
      24-JUL-1987; FR-010589.
      (INSP) Inst Pasteur.
 PA
 PI
      Danchin A, Glaser P, Ullmann A.
      WPI; 89-079098/11.
 DR
 DR
      N-PSDB; N90659.
      Cloning and expressing genes in multi-protein function system - by
PT
      transforming host cell with indicator gene producing protein which
PT
      interacts with expression prod. of the gene being cloned.
PT
      Claim 14; Fig 2(a-f); 24pp; French..
PS
CC
      Sequences encoding at least a part of AC able to hybridise with a
CC
      gene expressing a protein with AC activity, and proteins with AC
CC
      activity able to form immune complexes with specific antibodies, are
CC
      claimed. The isolated AC gene can be used to produce Bordetella
CC
      pertussis toxin.
SQ
      Sequence 1522 AA;
S0
      206A; 95 R; 55 N; 121D; O B; 8 C; 66 Q; 75 E; O Z; 190G; 32 H;
      61 I; 123L; 41 K; 24 M; 36 F; 40 P; 93 S; 86 T; 13 W; 36 Y; 121V;
20
                        9 Optimized Score
                                                       Significance =
Initial Score
                                                                       3.89
                                                   57
                                                       Mismatches
                                                                        258
Residue Identity =
                      19% Matches
                                                   71
Gaps
                       35
                           Conservative Substitutions
                                                                          0
                                20
              X
                                                     40
                                                               50
                                                                         60
                      10
                                          30
              MTAASMGPVRVAFVVLLALCSRPAVG@NCSGPCRCPDEPAPRCPAGVSLVLDGCGCCRVCAK
    KRTENVØYRHVELARVGØV-VEVDTLEHVØHIIGGAGNDSITGNAHD----NFLAGGSGDDRLDGGAGNDTL
                                              1060
                                   1050
    1020
                                                            1070
                                                                      1080
              1030
                         1040
                                                                         130
          70
                                           100
                                                               120
                    80
                                 90
                                                     1 14
    @LGELCTERDPCDPHKGLFCDFG==-SPANRKIGVCTAKDGAFCIFGGTVYKSGESF@SSCKY@CTCLDGAV
      1 1
```

```
1090
                 1100
                                              1130
                          1110
                                     1120
                                                             1140
             140
                       150
                                                                    190
                                    160
                                             170
                                                         180
    GCMP---LCSMDVRLPSPDCPFPR---RVKLPGKCCEEWVCDEPKDØTVVG--PALAAYRLEDT-FGPDPTM
    EPWPALNLFSVDHVKNIENLHGSRLNDRIAGDD@DNELWGHDGNDTIRGRGGDDILRGGLGLDTLYGEDGND
                         1180 1190
                                                           1210
          1160
                    1170
                                                 1200
                                                                     1220
       200
                210
                          220
                                    230
                                              240
                                                        250
                                                                  260
    IRANCLV@TTEWSACSKTCGMGISTRVTNDNASCRLEK@SRLCMVRPCEADLEENIKKGKKCIRTPKISKPI
    I----FL@DDETVSDDIDGGAGLHT-VDYSAMIHPGRIVAPHEYGFGIEADLSREWVRKASALGVDYYDNVR
            1230 1240 1250
                                                    1270
                                          1260
                                                              1280
     270
                  280
                           290
                                     300
                                               310
                                                         320
                                                                   330
    KFELSGCTSMK---TYRAKFCGVCTDGRCCTPHRTTTLPVEFKCPDGEVMKKNMMFIKTCACHYNCPGDNDI
           1 1 1 1
                                                NVENVIGTSMKDVLIGDA@ANTLMG@GGDDTGAAATANDLLF-GGDGNDMLYGDAGNDT---LYGGLGD-DT
 1290
           1300
                    1310 1320
                                        1330 1340
                                                                1350
      340
                X
    FESLYYRKMYGDMA
    LEGGAGNDWFG@T@AREHDVLRGG
                       1380
    1360
             1370
8. US-07-752-427-2 (1-349)
   R12108
               N-terminal deleted adenyl cyclase from Bacillus sp
     R12108 standard; protein; 1445 AA.
 ID
     R12108;
 AC
 DT
     30-JUL-1991 (first entry)
     N-terminal deleted adenyl cyclase from Bacillus sp.
 DE
 KW
     cyclic adenosine monophosphate; cAMP; NDFK.
 05
      Bacillus pertussis.
     WO9106664-A.
 PN
 PD
      16-MAY-1991.
 PF
     31-DCT-1990; F00793.
 PR
      31-0CT-1989; FR-014328.
 PR
      12-APR-1990; FR-004754.
 PA
      (INSP ) INST PASTEUR.
 PI
      Lacombe ML, Veron M, Mock M, Barzu O, Sarfati R;
 DR
     WPI; 91-164211/22.
 DR
     N-PSDB; @11826.
 PT
     Partly new substd. adenosine di: and tri:phosphate derivs. prodn.
 PT
     - useful as laboratory reagents e.g. affinity chromatography
 PT
      ligands or fluorescence markers
 PS
     Claim 11; Fig 1; 54pp; French.
 CC
      This protein is an N-terminal deletion of the protein encoded by
 CC
     B. pertussis "cya" gene. The first 261 N-terminal amino acids have
 CC
     been deleted, without affecting the adenyl cyclase activity of the
 CC
     protein. The truncated adenyl cyclase was obtained by unidirectional
CC
     deletion of the full-length cya gene, followed by expression in
     protease-deficient E. coli. The truncated enzyme is useful for
 CC
 CC
      synthesis of cAMP and adenosine phosphate derivatives; to separate
CC
     chemically synthesised 3'-substituted ATP derivatives and as a
CC
     reagent for assay of pyrophosphate, cAMP and calmodulin.
CC
     See also 011825 and W09106671-A.
50
                1445 AA;
     Sequence
SØ
     188A; 76 R; 58 N; 142D; O B; O C; 63 Q; 74 E; O Z; 214G; 27 H;
50
     58 I; 126L; 31 K; 17 M; 34 F; 31 P; 77 S; 67 T; 13 W; 34 Y; 115V;
                       9 Optimized Score
Initial Score
                                                 56 Significance =
                                                                     3.57
Residue Identity =
                     19%
                          Matches
                                                                      259
```

Mismatches

==

70

VORFRRIALATORARIO DO LEGITO DA PARENTA DE LA LA LA CALLA DE LA CALCA DEL CALCA DE LA CALCA DE LA CALCA DE LA CALCA DE LA CALCA DEL CALCA DE LA CALCA DEL CALCACA DEL CALCA DEL CALCA DEL CALCA DEL CALCA DEL CALCA DEL CALCA D

```
Gaps
                                 20
                                           30
                                                     40
                                                                50
                                                                          60
              X
                      10
              MTAASMGPVRVAFVVLLALCSRPAVGQNCSGPCRCPDEPAPRCPAGVSLVLDGCGCCRVCAK
    KRTENVØYRHVELARVGØV-VEVDTLEHVØHIIGGAGNDSITGNAHD----NFLAGGSGDDRLDGGAGNDTL
                                                                   780
                                750
                                          760
          730 X
                    740
                                                        770
                                                     110
                                                                120
                                  90
                                           100
          70
                    80
                                                                          130
    @LGELCTERDPCDPHKGLFCDFGSPAN----RKIGVCTAKDGAPCIFGGTVYRSGESF@SSCKY@CTCLDGAV
      1 1
    VGGEG@NTVIGGAGDDVFL@DLGVWSN@LDGGAGVDTVKYNVH@PSEERLERMG----DTGIHADL@KGTV
             800
                                  820
                                            830
                                                      840
                                                                      850
                       810
   790
                                                                       190
             140
                       150
                                     160
                                               170
                                                            180
    GCMP---LCSMDVRLPSPDCPFPR---RVKLPGKCCEEWVCDEPKD@TVVG--PALAAYRLEDT-FGPDPTM
    EKWPALNLFSVDHVKNIENLHGSRLNDRIAGDD@DNELWGHDGNDTIRGRGGDDILRGGLGLDTLYGEDGND
                                               900
                                                         910
                                                                    920
                870
                          880
                                     890
      860
                                                                     260
                                                240
                                                           250
                 210
                           220
                                      230
       200
    IRANCLVØTTEWSACSKTCGMGISTRVTNDNASCRLEKØSRLCMVRPCEADLEENIKKGKKCIRTPKISKPI
    I----FL@DDETVSDDIDGGAGLDT-VDYSAMIHPGRIVAPHEYGFGIEADLSREWVRKASALGVDYYDNVR
                                                  970
                                                             980
                  940
                                        960
                                                                       990
        930
                              950
     270
                            290
                                                 310
                                                            320
                                                                      330
                                       300
                  280.
    KFELSGCTSMK---TYRAKFCGVCTDGRCCTPHRTTTLPVEFKCPDGEVMKKNMMFIKTCACHYNCPGDNDI
    NVENVIGTSMKDVLIGDAQANTLMGQGGDDT-VRGGDGDDLLFGGDGNDMLYGDAGNDT---LYGGLGD-DT
                                      1030
                                                1040
                                                           1050
                1010
                          1020
                                                                         1060
      1000
      340
    FESLYYRKMYGDMA
    LEGGAGNDWFG@T@AREHDVLRGG
         1070
                 X 1080
9. US-07-752-427-2 (1-349)
                Platelet derived growth factor (PDGF) receptor pro
   R08267
      R08267 standard; protein; 1089 AA.
 ID
 AC
      R08267;
      07-MAR-1991 (first entry)
 DT
      Platelet derived growth factor (PDGF) receptor protein.
 DE
      Atherosclerosis; fibrotic diseases.
 KW
 OS 
      Homo sapiens.
 PN
      WD9014425-A.
 PD
      29-NOV-1990.
 PF
      21-MAY-1990; U02849.
 PR
      22-MAY-1989; US-355018.
 PA
      (ZYMO-) ZYMOGENETICS INC.
 PI
      Kelly JD, Murray MJ;
 DR
      WPI; 90-375992/50.
 DR
      N-PSDB; @06869.
 PT
      DNA encoding platelet-derived growth factor - used to transform
 PT
      cells for culturing to detect PDG agonists and antagonists
 PS
      Claim 1; Fig 1; 30pp; English.
 CC
      Gene product may be expressed from a transformed cell. It has
 CC
      utility in dection of PDGF agonist and antagonist analogues, binding
 CC
      AA, AB and BB isoforms. PDGF agonists may be used to enhance wound
      healing, and antagonists may be used to block the effects of PDGF
 CC
 CC
      eg. in treatment of a perosclerosis or fibrotic
                                                        seases.
 SQ
                 1089 AA;
      Sequence
      54 A; 44 R; 44 N; 66 D; O B; 17 C; 25 Q; 94 E; O Z; 58 G; 25 H;
 20
```

Conservative Substitutions

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timized Score =
                                                   56
                                                         nificance =
Initial Score
                        6
                                                                       3.57
Residue Identity =
                                                       Mismatches
                                                                        251
                           Matches
                                                   72
                     19%
                           Conservative Substitutions
Gaps
                       44
                                                                          0
                                                      50
                                                                    60
         X 10
                       20
                                 30
                                            40
    MTAASMGPVRVAFVVL-LALCSRPAVG@NCSGPCRCPDEPAPRCPAGVSLVLDGCGCCRV----CAK@LGEL
               1 1 1 1
         MGTSHPAFLVLGCLLTGLSLILCQLSLPSILPNENEKVVQLNSSFSLRCFGESEVSWQYPMSEEESS
                                     30
                                                40
         X
                 10
                           20
                                  100
                                             110
                                                           120
                                                                     130
     70
                         30
               80
    CTERDPCDPHKGLFCDFGSPANRKIGVCTAKDGAPCIFGGTVY----RSGESF@SSCKY@CTCLDGAVGCMP
    DVEIRNEENNSGLF-----VTLLEVSSA--SAAHTGLYTCYYNHT@TEENELEGRHIYIYVPDPDVAFVP
                                                      110
                                                                           130
                                90
                                            100
                                                                120
     70
               80
                                       170
      140
                             160
                                                  180
                                                              190
                   150
    LCSMDVRLPSPD---CPFPRRVKLPGKCCEEWVCDEPKD@TVVGPALAAYR--LEDTFGPDPTMIRA---NC
    LGMTDYLVIVEDDDSAIIPCRTTDP----ETPV--TLHNSEGVVPASYDSR@GFNGTFTVGPYICEATVKGK
                     150
                                                170
                                                          180
                                                                    190
           140
                                      160
  200
              210
                        220
                                  230
                                            240
                                                       250
                                                                 260
    LVQT--TEVSACSKTCGMGISTRVTNDNASCRLEKQSRLCMVRPCEADLEENIKKGKKCIRTPKISKPIKFE
   KF@TIPFNVYALKAT----SELDLEMEALKTVYK-SGETIVVTCAVFNNEVVDL@WTYPGEVKGKGITILE
                                                    240
                                                              250
     200
               210
                              220
                                         230
                                                                        260
                                               310
  270
                                   300
                                                                   330
               280
                         290
                                                         320
    LSGCTSMK---TYRAKFCGVCTDGRCCTPHRTTTLPV-EFKCPDGEVMKKNMMFIKTCACHYNCPGDNDIFE
    EIKVPSIKLVYTLTVPEATVKDSGDYECAARGATREVKEMKKVTISVHEKGFIEIKPTFSGLEAVNLHEVKH
                   280
                             290
                                       300
                                                  310
                                                            320
                                                                      330
         270
    340
    SLYYRKMYGDMA
    FVVEVRAYPPPRISWLKNNLTL
               X 350
       340
10. US-07-752-427-2 (1-349)
                 Alpha type PDGF receptor deduced from TR4 cDNA clo
    R06910
      R06910 standard; protein; 1089 AA.
 ID
AC
     R06910;
     16-JAN-1991 (first entry)
DT
     Alpha type PDGF receptor deduced from TR4 cDNA clone.
DE
     Platelet derived growth factor; T11.
KW
08
     Homo sapiens.
FH
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                      Location/Qualifiers
FT
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FT
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     /label=ligand binding domain
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     /label=transmembrane region
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     /label=tyrosine autophosphorylation site
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      07-SEP-1990.
 PF
      08-FEB-1990; U00617.
 PR
      09-FEB-1989; US-308282.
 PA
      (USDC ) US SEC OF COMMERCE.
 PI
      Matsui T, Aaronson SA, Pierce JH;
 DR
      WPI; 90-290306/38.
 DR
      N-PSDB; @05989.
 PT
      Type alpha platelet-derived growth factor receptor gene - useful
 PT
      for transforming cells to express novel protein receptor and also
 PT
      susceptible to genetic engineering.
 PS
      Claim 7; Fig 3; 64pp; English.
      The TR4 clone is the largest cDNA clone related to the T11 genomic
 CC
      clone, isolated from a library prepd. from human thymus DNA.
 CC
 CC
      T4 cDNA clone was isolated from a M426 human embryo fibroblast
      cDNA library. The coding region can be introduced into the pSV2
 CC
 CC
      gpt vector with a simian sarcoma virus LTR as a promoter and
 CC
      expressed in a host. The resulting protein is a novel PDGF
 CC
      receptor designated type alpha (the known receptor is designated
 CC
      type beta). The polypeptide has a calculated molecular mass of 120
 CC
      kD and has all the characteristics of a membrane spanning tyrosine
      kinase receptor. The extracellular region comprises a hydrophobic
 CC
      signal peptide and a ligand binding domain which has structural
 CC
      homology with the PDGF-R/CSF1-R subfamily. Ten Cys residues are
 CC
      spaced at the same positions as in other receptors of the sub-
 CC
 CC
      family and eight potential N-linked glycosylation sites are also
 CC
      present. A hydrophobic segment spans the membrane and the cyto-
 CC
      plasmic region comprises a juxtamembrane region, a tyrosine kinase
      region split into TK1 and TK2 by a hydrophilic interkinase region
 CC
      and a hydrophilic C-terminal tail. The TK region includes the
 CC
 CC
      consensus ATP binding sequence (G-X-G-X-X-G...K) and a tyrosine
 CC
      autophosphorylation site homologous to that of pp60(v-src).
SØ
                 1089 AA;
      Sequence
      54 A; 44 R; 44 N; 66 D; O B; 17 C; 26 Q; 93 E; O Z; 58 G; 25 H;
SD
SQ
      70 I; 104L; 69 K; 26 M; 32 F; 51 P; 93 S; 68 T; 13 W; 47 Y; 89 V;
Initial Score
                           Optimized Score
                                                  56 Significance =
                                                                       3. 57
                        6
Residue Identity =
                                                      Mismatches
                                                                        251
                           Matches
                                                  72
                      19%
                           Conservative Substitutions
Gaps
                       44
                                                                          0
         X 10
                       20
                                 30
                                           40
                                                     50
                                                                    60
   MTAASMGPVRVAFVVL-LALCSRPAVG@NCSGPCRCPDEPAPRCPAGVSLVLDGCGCCRV----CAK@LGEL
         MGTSHPAFLVLGCLLTGLSLILCQLSLPSILPNENEKVVQLNSSFSLRCFGESEVSW@YPMSEEESS
                 10
                                     30
                                                         50
         X
                           20
                                               40
                                                                    60
     70
                                  100
               80
                                            110
                                                           120
                                                                     130
                         30
   CTERDPCDPHKGLFCDFGSPANRKIGVCTAKDGAPCIFGGTVY----RSGESF@SSCKY@CTCLDGAVGCMP
               1 1 1
   DVEIRNEENNSGLF----
                        -VTVLEVSSA--SAAHTGLYTCYYNHT@TEENELEGRHIYIYVPDPDVAFVP
     70
                                                                120
                                                                          130
               80
                                90
                                                      110
                                           100
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